



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE
(Case No. 01-1720)

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9P-1614
P#4

In re Application of:)
Asp, et al.)
Serial No.: 09/993,211) Examiner: TBA
Filed: November 16, 2001) Art Unit: 1614
For: Method for the Treatment of)
Cardiotoxicity Induced by Antitumor)
Compounds)

TRANSMITTAL LETTER

Asst. Commissioner for Patents
Washington, D.C. 20231

Dear Sir:

In regard to the above identified application,

1. We are transmitting herewith the attached:
 - a) Supplemental Information Disclosure Statement;
 - b) PTO Form 1449 and 30 cited references;
 - c) Return postcard
2. With respect to fees:
 - a) It is believed no fee is due at this time.
 - b) Please charge any underpayment or credit any overpayment our Deposit Account, No. 13-2490.
3. GENERAL AUTHORIZATION: Please charge any additional fees or credit overpayment to Deposit Account No. 13-2490. A duplicate copy of this sheet is enclosed.
4. CERTIFICATE OF MAILING UNDER 37 CFR § 1.8: The undersigned hereby certifies that this Transmittal Letter and the paper, as described in paragraph 1, are being deposited with the United States Postal Service with sufficient postage as first class mail in an envelope addressed to the Asst. Commissioner for Patents, Washington, D.C. 20231 on June 5, 2002.

Date: June 5, 2002

Respectfully submitted,


Lisa M.W. Hillman

Registration No. 43,673



UNITED STATES PATENT AND TRADEMARK OFFICE
(Case No. 01-1720)

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In the Application of:

Asp, et al.

Serial No.: **09/993,211**

Filed: **November 16, 2001**

For: **Method for the Treatment of
Cardiotoxicity Induced by Antitumor
Compounds**

Art Unit: 1614

Examiner: To be assigned

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
Washington, D.C. 20231

Dear Sir:

Pursuant to the duty of disclosure provided by 35 C.F.R. § 1.56 and §§ 1.97-98, the applicants wish to make the following references of record in the above-identified application. Copies of the references are enclosed. Copies are also listed in the PTO-1449 form enclosed herewith. It is requested that the documents be given careful consideration and that they be cited of record in the prosecution history of the present application so that they will appear on the face of the patent issuing from the present application.

In the judgment of the undersigned, portions of the references may be material to the examination of the pending claims, however no such admission is intended. 37 C.F.R. 1.97 (h). The references have not been reviewed in sufficient detail to make any

other representation and, in particular, no representation is indented as to the relative importance of any portion of the references. This Statement is not a representation that the cited references have effective dates early enough to be "prior art" within the meaning of 35 U.S.C. sections 102 or 103.

CITED REFERENCES

Other Documents

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- 2) Lopez, et al., "*Randomized Prospective Clinical Trial of High-Dose Epirubicin and Dexrazoxane in Patients with Advanced Breast Cancer and Soft Tissue Sarcomas*", *J. Clin. Oncol.* Vol. 16, No. 1, pp. 86-92, 1998
- 3) Vici, et al., "*Dexrazoxane cardioprotection in advanced breast cancer patients undergoing high-dose epirubicin treatment*", *Clin. Ter.*, 149, pp. 15-20, 1998
- 4) Imondi, "*Preclinical Models of Cardiac Protection and Testing for Effects of Dexrazoxane on Doxorubicin Antitumor Effects*", *Semin. Oncol.* Vol. 25, No. 4, Suppl. 10, pp. 22-30, 1998
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- 9) Wexler, "*Ameliorating Anthracycline Cardiotoxicity in Children with Cancer: Clinical Trials with Dexrazoxane*", *Semin. Oncol.*, Vol. 25, No. 4, Suppl. 10, pp. 86-92, 1998
- 10) Synold, et al., "*Antineoplastic Activity of Continuous Exposure to Dexrazoxane: Potential New Role as a Novel Topoisomerase II Inhibitor*", *Semin. Oncol.*, Vol.

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- 12) Della Torre, et al., "*Protection against doxorubicin-induced cardiotoxicity in weanling rats by dexrazoxane*", *Cancer Chemother Pharmacol*, 43, pp. 151-156, 1999
- 13) O'Donnell, et al., "*A Phase I Trial of the VEGF Inhibitor SU5416, Incorporating Dynamic Contrast MRI Assessment of Vascular Permeability*", *685, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 177a, 2000
- 14) Baselga, et al., "*Continuous Administration of ZD1839 (Iressa), a Novel Oral Epidermal Growth Factor Receptor Tyrosine Kinase Inhibitor (EGFR-TKI), in Patients with Five Selected Tumor Types: Evidence of Activity and Good Tolerability*", *686, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 177a, 2000
- 15) Stewart, et al., "*Population Pharmacokinetics of Topotecan in Children and Adolescents*", *687, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 177a, 2000
- 16) Lewis, et al., "*A Phase-I Study of the Novel Topoisomerase-I Inhibitor J-107088 Administered on a Multiple-Dose Schedule*", *688, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 177a, 2000
- 17) Garrison, et al., "*A Phase I and Pharmacokinetic Study of the Camptothecin (CPT) Analog DX-89511 (Exetecan Mesylate): Escalating Infusion Duration and Dose*", *765, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 197a, 2000
- 18) Houghton, et al., "*Anticancer Activity of the Novel DB-67 Silatecan, a Highly Lipophilic Liposomal Camptothecin Agent Displaying Improved Intrinsic Human Blood and Plasma Stability and Potent Topoisomerase I Activity*", *766, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 197a, 2000
- 19) Peck, "*Phase I Trial of J-107088, a Novel Topoisomerase I Inhibitor, Administered Once Every 21 Days*", *767, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 197a, 2000
- 20) Bos, et al., "*Phase I Study of NX 211 (Liposomal Lurtotecan) Administered as a Single Dose Every 3 Weeks*", *768, *Proceedings of ASCO, Clinical Pharmacology*, Volume 19, p. 197a, 2000

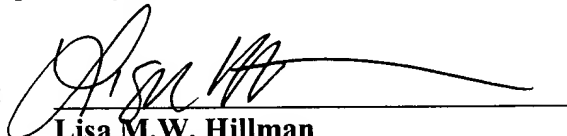
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- 22) Gianni, "*Tolerability in patients receiving trastuzumab with or without chemotherapy*", *Annals of Oncology*, 12 (Suppl. 1): S63-S68, 2001
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- 25) Malisza, et al., "*Inhibition of Anthracycline Semiquinone Formation by ICRF-187 (Dexrazoxane) in cells*", *Free Radical Biology & Medicine*, Vol. 20, No. 7, pp. 905-914, 1996
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- 27) Woodlock, et al., "*Coincident Acute Myelogenous Leukemia and Ischemic Heart Disease: Use of the Cardioprotectant Dexrazoxane During Induction Chemotherapy*", *American Journal of Hematology*, 59, pp. 246-248 1998
- 28) Sawyer, et al., "*Daunorubicin-Induced Apoptosis in Rat Cardiac Myocytes is Inhibited by Dexrazoxane*", *Original Contributions*, pp.257-265, © American Heart Association, Inc. 1999
- 29) Ohkubo, et al., "*Synthesis and Biological Activities of NB-506 Analogues Modified at the Glucose Group*", *Bioorganic & Medicinal Chemistry Letters*, Vol. 10, pp. 419-422, 2000

- 30) Long, et al., "*Non-camptothecin topoisomerase I active compounds as potential anticancer agents*", *Expert Opinion on Therapeutic Patents*, 10(5), pp. 635-666, 2000

Respectfully submitted,

Date: June 5, 2002

by:

A handwritten signature in dark ink, appearing to read 'Lisa M.W. Hillman', written over a horizontal line.

Lisa M.W. Hillman

Reg. No. 43,673

8	Hellmann, "Overview and Historical Development of Dexrazoxane", <i>Semin. Oncol.</i> , Vol. 25, No. 4, Suppl. 10, pp. 48-54, 1998
9	Wexler, "Ameliorating Anthracycline Cardiotoxicity in Children with Cancer: Clinical Trials with Dexrazoxane", <i>Semin. Oncol.</i> , Vol. 25, No. 4, Suppl. 10, pp. 86-92, 1998
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22	Gianni, "Tolerability in patients receiving trastuzumab with or without chemotherapy", <i>Annals of Oncology</i> , 12 (Suppl. 1): S63-S68, 2001
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29	Ohkubo, et al., "Synthesis and Biological Activities of NB-506 Analogues Modified at the Glucose Group", <i>Bioorganic & Medicinal Chemistry Letters</i> , Vol. 10, pp. 419-422, 2000
30	Long, et al., "Non-camptothecin topoisomerase I active compounds as potential anticancer agents", <i>Expert Opinion on Therapeutic Patents</i> , 10(5), pp. 635-666, 2000

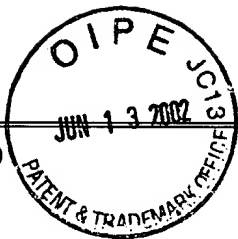
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FORM PTO-1449
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Atty. Docket N .

01-1720

S rial No.

09/993,211

INFORMATION DISCLOSURE
STATEMENT BY APPLICANT
(Use several sheets if necessary)

Applicant:

Asp, et al.

Filing Date:

November 16, 2001

Group:

1614

U.S. PATENT DOCUMENTS

Examiner Initial	Document Number	Date	Name	Class	Subclass	Filing Date if Appropriate

FOREIGN PATENT DOCUMENTS

	Document Number	Date	Country	Class	Subclass	Translation	
						Yes	No

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc).

1	Hasinoff, et al., "Chemical, Biological and Clinical Aspects of Dexrazoxane and Other Bisdioxopiperazines", <i>Current Medicinal Chemistry</i> , 5, p. 1-28, 1998
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